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## PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

* * *	* *	* *	* *	* Welcome to STN International
NEWS				Web Page for STN Seminar Schedule - N. America
NEWS	2	NOV	21	CAS patent coverage to include exemplified prophetic substances identified in English-, French-, German-, and Japanese-language basic patents from 2004-present
NEWS	3	NOV	26	MARPAT enhanced with FSORT command
NEWS	4	NOV		CHEMSAFE now available on STN Easy
	-			
NEWS	5	NOV		Two new SET commands increase convenience of STN searching
NEWS	6	DEC	01	ChemPort single article sales feature unavailable
NEWS	7	DEC	12	GBFULL now offers single source for full-text coverage of complete UK patent families
NEWS	8	DEC	17	Fifty-one pharmaceutical ingredients added to PS
NEWS	9	JAN		The retention policy for unread STNmail messages
NEWS	-	JAN		will change in 2009 for STN-Columbus and STN-Tokyo WPIDS, WPINDEX, and WPIX enhanced Japanese Patent
			-	Classification Data
NEWS	11	FEB	02	Simultaneous left and right truncation (SLART) added for CERAB, COMPUAB, ELCOM, and SOLIDSTATE
NEWS	12	FEB	02	GENBANK enhanced with SET PLURALS and SET SPELLING
NEWS	13	FEB	06	Patent sequence location (PSL) data added to USGENE
NEWS				COMPENDEX reloaded and enhanced
NEWS		FEB		WTEXTILES reloaded and enhanced
NEWS	16	FEB	19	New patent-examiner citations in 300,000 CA/CAplus patent records provide insights into related prior art
NEWS	17	FEB	19	Increase the precision of your patent queries use terms from the IPC Thesaurus, Version 2009.01
NEWS	18	FEB	23	Several formats for image display and print options discontinued in USPATFULL and USPAT2
NEWS	19	FEB	23	MEDLINE now offers more precise author group fields and 2009 MeSH terms
NEWS	20	FEB	23	TOXCENTER updates mirror those of MEDLINE - more precise author group fields and 2009 MeSH terms
NEWS	21	FEB	23	Three million new patent records blast AEROSPACE into STN patent clusters
NEWS	22	FEB	25	USGENE enhanced with patent family and legal status display data from INPADOCDB
NEWS	23	MAR	06	INPADOCDB and INPAFAMDB enhanced with new display formats
NEWS	24	MAR	11	EPFULL backfile enhanced with additional full-text
	0.5			applications and grants
NEWS		MAR		ESBIOBASE reloaded and enhanced
NEWS	26	MAR	20	CAS databases on STN enhanced with new super role

for nanomaterial substances

NEWS 27 MAR 23 CA/Caplus enhanced with more than 250,000 patent equivalents from China

NEWS 28 MAR 30 IMSPATENTS reloaded and enhanced

NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3, AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

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NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8

Enter NEWS followed by the item number or name to see news on that specific topic.

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FILE 'HOME' ENTERED AT 09:37:25 ON 30 MAR 2009

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FILE 'REGISTRY' ENTERED AT 09:37:49 ON 30 MAR 2009 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2009 American Chemical Society (ACS)

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=> file caplus
COST IN U.S. DOLLARS SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST 0.48 0.70

FILE 'CAPLUS' ENTERED AT 09:38:06 ON 30 MAR 2009
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FILE COVERS 1907 - 30 Mar 2009 VOL 150 ISS 14 FILE LAST UPDATED: 29 Mar 2009 (20090329/ED)

Caplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2008.

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## http://www.cas.org/legal/infopolicy.html

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=> s 11 SSS full

REGISTRY INITIATED

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FULL SEARCH INITIATED 09:38:09 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 991122 TO ITERATE

99.2% PROCESSED 983210 ITERATIONS

99 ANSWERS

100.0% PROCESSED 991122 ITERATIONS

99 ANSWERS

SEARCH TIME: 00.00.26

1.2 99 SEA SSS FUL L1

L3 11 L2

AUTHOR(S):

SOURCE:

CORPORATE SOURCE:

=> d ibib abs hitstr 1-

YOU HAVE REQUESTED DATA FROM 11 ANSWERS - CONTINUE? Y/(N):v

ANSWER 1 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2006:625349 CAPLUS Full-text

DOCUMENT NUMBER: 145:224321

TITLE: The synthesis and SAR of

2-arylsulfanylphenyl-1-oxyalkylamino acids as GlyT-1 inhibitors

Smith, Garrick; Mikkelsen, Gitte; Eskildsen, Jorgen;

Bundgaard, Christoffer

Medicinal Chemistry Research, H. Lundbeck A/S, Valby,

DK 2500, Den.

Bioorganic & Medicinal Chemistry Letters (2006), 16(15), 3981-3984

CODEN: BMCLE8; ISSN: 0960-894X

Elsevier B.V.

PUBLISHER: DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 145:224321 GI

- AB Elevation of glycine levels by inhibition of the glycine transporter-1 [GlyT-1] and activation of the NMDA receptor is a potential strategy for the treatment of schizophrenia. A novel series of 2-arylsulfanylphenyl-1-oxyalkyl amino acids have been identified. The most prominent member of this series [I] is a potent GlyT-1 inhibitor (ICSO = 59 nM). In vitro and in vivo assessment of CNS exposure indicates this compound is a likely substrate for active efflux transporters.
- TT 791643-06-6P 793643-10-2P 791643-25-9P 791643-27-1P 791643-27-1P 791643-27-1P 791643-31-7P 791643-68-0P 905815-55-4P 905815-55-4P 905815-55-4P 905815-55-4P 905815-55-4P 905815-59-0P 905815-690-3P 905815-61-4P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(synthesis and SAR of arylsulfanylphenyloxyalkylamino acids as GlyT-1 inhibitors)

RN 791643-06-6 CAPLUS

CN Glycine, N-[2-[2-[(3-chlorophenyl)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

RN 791643-10-2 CAPLUS

CN Glycine, N-[2-[2-[(3-fluorophenyl)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

RN 791643-25-9 CAPLUS

CN Glycine, N-[2-[2-[(3-chloropheny1)thio]phenoxy]ethyl]-N-ethyl- (CA INDEX NAME)

RN 791643-27-1 CAPLUS

CN Glycine, N-[2-[2-[(4-chlorophenyl)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

RN 791643-31-7 CAPLUS

CN Glycine, N-[2-[2-[[4-(1,1-dimethylethyl)phenyl]thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

RN 791643-68-0 CAPLUS

CN Glycine, N-[(1R)-2-[2-[(3-fluoropheny1)thio]phenoxy]-1-methylethyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 905815-53-4 CAPLUS

CN Glycine, N-[2-[2-[(4-methoxyphenyl)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

RN 905815-54-5 CAPLUS

CN Glycine, N-[2-[2-[(4-fluorophenyl)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

RN 905815-55-6 CAPLUS

CN Glycine, N-methyl-N-[2-[2-[(4-methylphenyl)thio]phenoxy]ethyl]- (CA INDEX NAME)

RN 905815-56-7 CAPLUS

CN Glycine, N-[2-[2-[[4-(acetylamino)phenyl]thio]phenoxy]ethyl]-N-methyl-(CA INDEX NAME)

RN 905815-57-8 CAPLUS

CN Glycine, N-[1-[[2-[(3-fluoropheny1)thio]phenoxy]methyl]propyl]-N-methyl-(CA INDEX NAME)

RN 905815-58-9 CAPLUS

CN Glycine, N-[1-[[2-[(3-fluorophenyl)thio]phenoxy]methyl]-2-methylpropyl]-N-methyl- (CA INDEX NAME)

RN 905815-59-0 CAPLUS

CN Glycine, N-[(1S)-2-[2-[(3-fluorophenyl)thio]phenoxy]-1-methylethyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 905815-60-3 CAPLUS

CN L-Alanine, N-[2-[2-[(3-chlorophenyl)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 905815-61-4 CAPLUS

CN L-Alanine, N-[2-[2-[(3-chlorophenyl)thio]phenoxy]ethyl]-N-ethyl- (CA INDEX NAME)

Absolute stereochemistry.

REFERENCE COUNT: 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 2 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:1154515 CAPLUS Full-text

ACCESSION NUMBER: 2005:11545 DOCUMENT NUMBER: 143:422634

TITLE: Preparation of N-(2-aryloxyethyl)glycine derivatives

and their use as glycine transport inhibitors
INVENTOR(S): Man, Teresa; Milot, Guy; Porter, Warren Jaye; Reel,

Jon Kevin; Rudyk, Helene Catherine Eugenie; Valli,

Matthew John; Walter, Magnus Wilhelm PATENT ASSIGNEE(S): Eli Lilly and Company, USA

SOURCE: PCT Int. Appl., 187 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: English

LANGUAGE: Englis

PATENT INFORMATION:

PA:	PATENT NO.				KIN	D	DATE			APPL	ICAT		D					
						-												
WO	2005	1003	01		A1 20051027					WO 2	005-	JS89	62		20050318			
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		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FΙ,	GB,	GD,	
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	ΚZ,	LC,	
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,	
		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SM,	
		SY,	TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	
		AZ,	BY,	KG,	KΖ,	MD,	RU,	ΤJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	
		EE,	ES,	FΙ,	FR,	GB,	GR,	HU,	IE,	IS,	IT,	LT,	LU,	MC,	NL,	PL,	PT,	

RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,

MR, NE, SN, TD, TG

US 2004-558260P

PRIORITY APPLN. INFO.: OTHER SOURCE(S):

CASREACT 143:422634; MARPAT 143:422634

Z A NOTE OH

AB The invention relates to (aryloxyethyl)glycine derivs. I [X is H, halo, alkyl, CF3, cycloalkyl, aryloarbonyl, (un) substituted aryl, fused arylcycloalkyl or heteroaryl, fused arylheterocyclyl; Z is alkyl, alkenyl, Y, CO-Y, CH(OH)-Y, OY, alkyl-Y, alkyl-OY, SY, CF2Y or NRZ-Y, where Y is alkyl, (CH2)1-10CF3, CF3, CF3, C3F7, (un) substituted aryl, heteroaryl, cycloalkyl or heterocyclyl and R2 is H or alkyl; A is (un) substituted aryl, H, alkoxy; R1 is alkyl] or their pharmaceutically-acceptable salts that exhibit activity as inhibitors of the glycine type-1 transporter, to pharmaceutical compns. containing them and to their use in the treatment of neurol and neuropsychiatric disorders. Thus, glycine derivative I (X = Ph, Z = 2-thienyl, A = H, R1 = H) was prepared via reactions of 3-iodo-4-methoxybiphenyl, 2-thiopheneboronic acid, and [(2-hydroxyethyl)methylaminolacetic acid text-Bu ester.

IT 868263-52-9P 868264-97-5P 868265-44-5P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of (aryloxyethyl)glycine derivs. as glycine transport inhibitors)

RN 868263-52-9 CAPLUS

CN Glycine, N-methyl-N-[2-[(3-phenoxy[1,1'-biphenyl]-4-yl)oxy]ethyl]- (CA INDEX NAME)

RN 868264-97-5 CAPLUS

CN Glycine, N-[2-[[3-(cyclohexyloxy)[1,1'-biphenyl]-4-yl]oxy]ethyl]-N-methyl-(CA INDEX NAME)

RN 868265-44-5 CAPLUS

CN Glycine, N-methyl-N-[2-[[3-(phenylthio)[1,1'-biphenyl]-4-yl]oxy]ethyl]-(CA INDEX NAME)

IT 868263-20-1P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of (aryloxyethyl)glycine derivs. as glycine transport inhibitors)

RN 868263-20-1 CAPLUS

CN Glycine, N-methyl-N-[2-[[3-(phenylthio)[1,1'-biphenyl]-4-yl]oxy]ethyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 3 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:965214 CAPLUS Full-text

DOCUMENT NUMBER: 141:411217

TITLE: A preparation of oxyphenyl and sulfanylphenyl

derivatives of amino acids, useful as glycine

transporter inhibitors

INVENTOR(S): Smith, Garrick Paul; Mikkelsen, Gitte; Andersen, Kim;

Greve, Daniel Rodriguez; Eskildsen, Joergen

PATENT ASSIGNEE(S): H. Lundbeck A/S, Den.

SOURCE: PCT Int. Appl., 87 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

AB

PA:	TENT	NO.			KIND DATE						ICAT								
WO	2004	0967	61													0040	427		
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,		
		CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	ΚZ,	LC,		
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,		
		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,		
		TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW		
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,		
		ΑZ,	BY,	KG,	KΖ,	MD,	RU,	ΤJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,		
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,		
		SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	ΝE,		
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$$R^3$$
 $R^4$ 
 $X$ 
 $R^7$ 
 $R^9$ 
 $R^{11}$ 
 $R^{10}$ 
 $R^{10}$ 

The invention relates to a preparation of aromatic oxyphenyl and aromatic sulfanylphenyl derivs. of formula I (wherein: X is 0, S, or CH2, etc.; Y is 0 or S; R1, R2, R3, and R4 are independently selected from H, halogen, CN, NO2, or alk(en/yn)yl, etc.; R5 is (un)substituted aryl or monocyclic heteroaryl; R6

is H, alk(en/yn)yl, cycloalk(en)yl, or alk(en/yn)ylsulfanyl, etc.; R7 and R8 are independently selected from H, alk(en/yn)yl, or cycloalk(en)yl; R9 and R11 are independently selected from H, alk(en/yn)yl, hydroxyalk(en/yn)yl, or alk(en/yn)ylsulfanyl, etc.; R10 is H, alk(en/yn)yl, aryl, or arylalk(en/yn)yl, etc.; R6 and R8 together with the nitrogen may form 3-7 membered heterocyclic ring], useful as glycine transporter inhibitors (IC50 < 10000 nM). The compds. of formula I are useful for the treatment of diseases such as schizophrenia, including both the pos. and the neg. symptoms of schizophrenia. For instance, pyrrolidinecarboxylic acid derivative II was prepared via etherification of 2-(3-fluorophenylsulfanyl)phenol by (hydroxyethyl)pyrrolidinecarboxylate derivative III.

791643-02-2P 791643-06-6P 791643-08-8P IΤ

791643-10-2P 791643-12-4P 791643-14-6P 791643-16-8P 791643-20-4P 791643-21-5P

791643-25-9P 791643-27-1P,

[[2-[2-(4-Chlorophenylsulfanyl)phenoxy]ethyl]-N-methylamino]acetic acid 791643-30-6P, [[2-[2-(4-tert-Butylphenylsulfanyl)phenoxy]ethyl]-N-

isopropylamino]acetic acid 791643-31-7P,

[[2-[2-(4-tert-Butvlphenvlsulfanvl)phenoxv]ethvl]-N-methvlamino]acetic acid 791643-33-9P, [[2-[2-(3,4-Dichlorophenylsulfanyl)phenoxy]ethyl]-N-methylamino]acetic acid

791643-34-0P 791643-45-3P,

[[1-[2-(3,4-Dichlorophenylsulfanyl)phenoxy]butan-2-yl]-N-ethylamino]acetic

acid 791643-46-4P 791643-48-6P 791643-49-7P

791643-51-1P 791643-52-2P 791643-53-3P

791643-55-5P 791643-57-7P 791643-58-8P

791643-63-5P 791643-65-7P 791643-66-8P

791643-68-0P 791643-70-4P 791643-71-5P 791643-72-6P 791643-73-7P 791643-74-8P

791643-75-9P 791643-76-0P 791643-77-1P

791643-78-2P 791643-79-3P 791643-80-6P

791643-81-7P 791643-84-0P 791643-86-2P 791643-87-3P 791644-10-5P 791644-11-6P

791644-12-7P 791644-16-1P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of oxyphenyl and sulfanylphenyl derivs. of amino acids, useful as glycine transporter inhibitors)

RN 791643-02-2 CAPLUS

CN Glycine, N-(2-(2-(4-(1.1-dimethylethyl)phenyl)thio)phenoxylethyl]-N-ethyl-(CA INDEX NAME)

791643-06-6 CAPLUS RN

CN Glycine, N-[2-[2-[(3-chloropheny1)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

- RN 791643-08-8 CAPLUS
- CN Glycine, N-[2-[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]ethyl]-N-methyl-(CA INDEX NAME)

- RN 791643-10-2 CAPLUS
- CN Glycine, N-[2-[2-[(3-fluorophenyl)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

- RN 791643-12-4 CAPLUS
- CN Glycine, N-(1-methylethyl)-N-[2-[2-[[4-

(trifluoromethyl)phenyl]thio]phenoxy]ethyl]- (CA INDEX NAME)

- RN 791643-14-6 CAPLUS
- CN Glycine, N-[2-[2-[(3,4-dichlorophenyl)thio]phenoxy]ethyl]-N-ethyl- (CA INDEX NAME)

RN 791643-16-8 CAPLUS

CN Glycine, N-ethyl-N-[2-[2-[[4-(methylthio)phenyl]thio]phenoxy]ethyl]- (CA INDEX NAME)

RN 791643-20-4 CAPLUS

CN Glycine, N-[2-[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]ethyl]-N-ethyl-(CA INDEX NAME)

RN 791643-21-5 CAPLUS

CN Glycine, N-(1-methylethyl)-N-[2-[2-[[4-(methylthio)phenyl]thio]phenoxy]ethyl]- (CA INDEX NAME)

RN 791643-25-9 CAPLUS

CN Glycine, N-[2-[2-[(3-chloropheny1)thio]phenoxy]ethyl]-N-ethyl- (CA INDEX NAME)

RN 791643-27-1 CAPLUS

CN Glycine, N-[2-[2-[(4-chlorophenyl)thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

RN 791643-30-6 CAPLUS

CN Glycine, N-[2-[2-[[4-(1,1-dimethylethyl)phenyl]thio]phenoxy]ethyl]-N-(1methylethyl)- (CA INDEX NAME)

RN 791643-31-7 CAPLUS

CN Glycine, N-[2-[2-[[4-(1,1-dimethylethyl)phenyl]thio]phenoxy]ethyl]-N-methyl- (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{Ho}_2\text{C-CH}_2 - \text{N-CH}_2 - \text{CH}_2 - \text{O} \\ \\ \end{array} \\ \text{S-CH}_2 \\ \text{Bu-t} \\ \end{array}$$

RN 791643-33-9 CAPLUS

CN Glycine, N-[2-[2-[(3,4-dichloropheny1)thio]phenoxy]ethy1]-N-methy1- (CA INDEX NAME)

- RN 791643-34-0 CAPLUS
- CN Glycine, N-methyl-N-[2-[2-[[4-(trifluoromethyl)phenyl]thio]phenoxy]ethyl]-(CA INDEX NAME)

- RN 791643-45-3 CAPLUS
- CN Glycine, N-[1-[[2-[(3,4-dichlorophenyl)thio]phenoxy]methyl]propyl]-N-ethyl-(CA INDEX NAME)

- RN 791643-46-4 CAPLUS
- CN Glycine, N-[1-[[2-[(3,4-dichlorophenyl)thio]phenoxy]methyl]-2methylpropyl]-N-ethyl- (CA INDEX NAME)

- RN 791643-48-6 CAPLUS
- CN Glycine, N-[1-[[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]methyl]-2methylpropyl]-N-ethyl- (CA INDEX NAME)

- RN 791643-49-7 CAPLUS
- CN Glycine, N-[2-[2-[(3-chlorophenyl)thio]phenoxy]-1-methylethyl]-N-ethyl-(CA INDEX NAME)

- RN 791643-51-1 CAPLUS
- CN Glycine, N-[1-[[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]methyl]butyl]-N-ethyl- (CA INDEX NAME)

- RN 791643-52-2 CAPLUS
- CN Glycine, N-[2-[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]-1-methylethyl]-N-ethyl- (CA INDEX NAME)

RN 791643-53-3 CAPLUS

CN Glycine, N-[(1S)-2-[2-[(3-chlorophenyl)thio]phenoxy]-1-methylethyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 791643-55-5 CAPLUS

CN Glycine, N-[(1S)-2-[2-[(3-chlorophenyl)thio]phenoxy]-1-methylethyl]-Nethyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 791643-57-7 CAPLUS

CN Glycine, N-[2-[2-[(3,4-dichlorophenyl)thio]phenoxy]-1-methylethyl]-N-ethyl-(CA INDEX NAME)

RN 791643-58-8 CAPLUS

CN Glycine, N-[2-[2-[(4-chlorophenyl)thio]phenoxy]-1-methylethyl]-N-ethyl-(CA INDEX NAME)

RN 791643-63-5 CAPLUS

CN Glycine, N-[(1R)-2-[2-[(3,4-dichlorophenyl)thio]phenoxy]-1-methylethyl]-N-ethyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 791643-65-7 CAPLUS
- CN Glycine, N-[(1R)-2-[2-(4-chlorophenoxy)phenoxy]-1-methylethyl]-N-methyl-(CA INDEX NAME)

Absolute stereochemistry.

- RN 791643-66-8 CAPLUS
- CN Glycine, N-[(1R)-2-[2-[(3-chlorophenyl)thio]phenoxy]-1-methylethyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 791643-68-0 CAPLUS

CN Glycine, N-[(1R)-2-[2-[(3-fluorophenyl)thio]phenoxy]-1-methylethyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 791643-70-4 CAPLUS

CN Glycine, N-[2-[2-[(3-chlorophenyl)thio]phenoxy]propyl]-N-ethyl- (CA INDEX NAME)

RN 791643-71-5 CAPLUS

CN Glycine, N-[1-[[2-[(3-chlorophenyl)thio]phenoxy]methyl]-2-methylpropyl]-Nmethyl- (CA INDEX NAME)

RN 791643-72-6 CAPLUS

CN Glycine, N-ethyl-N-[2-methyl-1-[[2-[[4-(trifluoromethyl)phenyl]thio]phenoxy]methyl]propyl]- (CA INDEX NAME)

RN 791643-73-7 CAPLUS

CN Glycine, N-[1-[[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]methyl]propyl]-N-methyl- (CA INDEX NAME)

RN 791643-74-8 CAPLUS

CN Glycine, N-[(1S)-2-[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]-1-methylethyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 791643-75-9 CAPLUS

CN Glycine, N-[(2S)-2-[2-[(3-fluorophenyl)thio]phenoxy]propyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 791643-76-0 CAPLUS

CN Glycine, N-[1-[[2-[[4-(1,1-dimethylethyl)phenyl]thio]phenoxy]methyl]-2-methylpropyl]-N-ethyl- (CA INDEX NAME)

RN 791643-77-1 CAPLUS

CN Glycine, N-[(1S)-2-[2-[(3,4-dichlorophenyl)thio]phenoxy]-1-methylethyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 791643-78-2 CAPLUS
- CN Glycine, N-[1-[[2-[(3-chloro-4-fluoropheny1)thio]phenoxy]methyl]-2methylpropyl]-N-methyl- (CA INDEX NAME)

RN 791643-79-3 CAPLUS

CN Glycine, N-[1-[[2-[[4-(1,1-dimethylethyl)phenyl]thio]phenoxy]methyl]propyl]-N-ethyl- (CA INDEX NAME)

RN 791643-80-6 CAPLUS

CN Glycine, N-[2-[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]propyl]-N-ethyl-(CA INDEX NAME)

RN 791643-81-7 CAPLUS

CN Glycine, N-cyclohexyl-N-[2-[2-[(4-methoxyphenyl)thio]phenoxy]propyl]- (CA INDEX NAME)

RN 791643-84-0 CAPLUS

CN Glycine, N-[2-[2-[(3-chlorophenyl)thio]phenoxy]propyl]-N-cyclohexyl- (CA INDEX NAME)

RN 791643-86-2 CAPLUS

CN L-Alanine, N-[2-[[3-[(3-fluorophenyl)thio][1,1'-biphenyl]-4-yl]oxy]ethyl]-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 791643-87-3 CAPLUS

CN Glycine, N-[2-[[3-[(3-fluorophenyl)thio][1,1'-biphenyl]-4-yl]oxy]ethyl]-N-methyl- (CA INDEX NAME)

RN 791644-10-5 CAPLUS

CN Glycine, N-[1-[[2-[(3-chloro-4-fluorophenyl)thio]phenoxy]methyl]propyl]-N-ethyl- (CA INDEX NAME)

RN 791644-11-6 CAPLUS

CN Glycine, N-[(2R)-2-[2-[(3-chlorophenyl)thio]phenoxy]propyl]-N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 791644-12-7 CAPLUS

CN Glycine, N-[2-[2-[(3-fluorophenyl)thio]phenoxy]propyl]-N-methyl- (CA INDEX NAME)

RN 791644-16-1 CAPLUS

CN L-Alanine, N-[2-[[3-[(3-fluorophenyl)thio][1,1'-biphenyl]-4-yl]oxy]ethyl]N-methyl- (CA INDEX NAME)

Absolute stereochemistry.

## REFERENCE COUNT: THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 4 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:878155 CAPLUS Full-text

DOCUMENT NUMBER: 141:366240

TITLE: Preparation of pyrrolopyridazines for the treatment of

proliferative disorders

INVENTOR(S): Salvati, Mark E.; Illig, Carl R.; Wilson, Kenneth Jerome; Chen, Jinsheng; Meegalla, Sanath K.; Wall,

Mark James

PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA

SOURCE: U.S. Pat. Appl. Publ., 189 pp., Cont.-in-part of U.S.

> Ser. No. 396,197. CODEN: USXXCO

DOCUMENT TYPE: Patent English

LANGUAGE: FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

		KIND DATE					APPL												
	2004						2004	1021							20030926				
	7030				B2		2006												
	2004									US 2	003-	3961	97	20030325					
	6900				B2		2005								_				
	2005				A2		2005			WO 2	004-	US31	571		20040923				
WO	2005						2005												
	W:						AU,												
							DE,												
							ID,												
							LV, PL,												
							TZ,												
	DW.						MW.												
							RU,												
							GR,												
							CF.												
			TD.		,	,	,	,	,	,	,	,		,	,	,	,		
EP	1664	051			A2		2006		EP 2	004-	7890		20040923						
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,		
		IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	PL,	SK,	HR	
US	2005	0159	420		A1		2005	0721		US 2	005-	2954	7		2	0050	105		
ORITY	APP	LN.	INFO	. :						US 2	003-	3961	97		A2 2	0030	325		
										US 2	002-	3682		P 20020328					
										US 2									
										US 2									
										WO 2	004-	US31	571		W 2	0040	923		
IER SC	ER SOURCE(S):					MARPAT 141:366240													

OTHER SOURCE(S): MARPAT 141:366240

GI

$$\begin{array}{c} R3 & YR4 \\ R2X & R1 \end{array}$$

AB Pyrrolopyridazines I [R1 = H, alkyl, aralkyl, halo, OH, etc.; R2 = alkyl, cycloalkyl, aryl, heterocyclic, aralkyl, (un)substituted CO2H, CHO, CONH2, SO3H, SO2NH2, SH, S(O)H, SO2H; R1R2, R2R3 = cycloalkyl, aryl, heterocyclic; R3 = H, alkyl, cycloalkyl, heterocyclic, aryl, aralkyl, acyl, halo, (un) substituted OH, CH2OH, CH2NH2, CH2SH; R4 = alkyl, cycloalkyl, aryl, heterocyclic, aralkyl, acyl, (un)substituted CO2H, CONH2, SO3H, SO2NH2, SH, S(O)H, SO2H; R5 = H, halo, CN, alkyl, cycloalkyl, heterocyclic, aryl, aralkyl, acyl, alkylene, (un) substituted CO2H, CONH2, SO3H, SO2NH2, SH, S(O)H, SO2H; R6 = H, alkyl, cycloalkyl, aryl, aralkyl, heterocyclic, acyl, alkoxycarbonyl, carbamoyl; X, Y, Z = bond, O, S, (un)substituted NH, etc.] were prepared for use in the treatment of proliferative, inflammatory, and other disorders (no data). Thus, NCCH2CO2Et was cyclized with MeCHO to di-Et 3-methyl-1H-pyrrole-2,4-dicarboxvlate which was N-aminated and cyclized with (EtO)2CHCH2CN to give Et 3-cyano-1,4-dihydro-5-methyl-4- oxopyrrolo[1,2-b]pyridazine-6-carboxylate. This ketone was chlorinated and treated with cyclohexylamine to give the title compound II. The compds. I were tested against several different kinases such as VEGFR-2, FGFR-1, HER-1, HER-2, HER-4, MEK and p38 kinases. Thus, tested compds. I inhibited VEGFR-2 and FGFR-1 kinases with IC50 of ≤80 µM.

775344-57-9P
RI: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of pyrrolopyridazines for the treatment of proliferative disorders)

RN 779344-57-9 CAPLUS

ΙT

CN

 $\label{eq:pyrolo} Pyrrolo[1,2-b] pyridazine-6-carboxylic acid, $3-cyano-4-[(4-[2-[2-[(2-(1,1-dimethylethoxy)-2-oxoethyl]amino]-1,1-dimethyl-2-oxoethoxy] phenoxy] phenyl] amino]-5-methyl-, methyl ester (CA INDEX NAME)$ 

IT 779344-58-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of pyrrolopyridazines for the treatment of proliferative disorders)

RN 779344-58-0 CAPLUS

CN Pyrrolo[1,2-b]pyridazine-6-carboxylic acid,

4-[[4-[2-[2-[(carboxymethyl)amino]-1,1-dimethyl-2-oxoethoxy]phenoxy]phenoxy]amino]-3-cyano-5-methyl-, 6-methyl ester (CA INDEX NAME)

REFERENCE COUNT: 30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 5 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2002:946033 CAPLUS Full-text

DOCUMENT NUMBER: 138:20910

TITLE: Preparation of

3-Methyl-2,6-dioxo-4-(trifluoromethyl)-1,2,3,6tetrahydropyrimidine derivatives as plant growth

regulators for cotton

INVENTOR(S): Mito, Nobuaki

PATENT ASSIGNEE(S): Sumitomo Chemical Company, Limited, Japan

SOURCE: PCT Int. Appl., 78 pp.

CODEN: PIXXD2 Patent

DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	TENT				KIND DATE						ICAT								
							2002	1212							20010531				
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,		
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,		
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	KE,	KG,	KR,	KZ,	LC,	LK,	LR,	LS,	LT,		
	LU, LV, MA,						MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PL,	PT,	RO,	RU,		
	SD, SE, SG,						SL,	ΤJ,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,		
	YU, ZA, ZW																		
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	AT,	BE,	CH,	CY,		
		DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,		
		ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG				
					A1 20021216					AU 2	001-		20010531						
					B2 200701														
										BR 2001-17032						0010	531		
US	2004	0152	597		A1		2004	0805		US 2	003-	4765		20031103					
US	7115	544			B2		2006	1003											
PRIORIT	Y APP	LN.	INFO	. :						WO 2	001-	JP45	84	1	vi 2	0010	531		
OTHER S	HER SOURCE(S):						MARPAT 138:20910												
GI																			

$$F3C \xrightarrow{\text{Me}} 0 \xrightarrow{R3} \xrightarrow{X} \xrightarrow{R2} \xrightarrow{R1}$$

AB Plant growth regulators for cotton containing as an active ingredient a compound I (X = CH, or N, Z = halo; A = O, S, or NH; RI = OH, CI-C7 alkoxy, C3-C7 alkenyloxy, C3-C7 alkynyloxy, C5-C7 cycloalkoxy, [di(C1-C7 alkoxy)carbonyl]C1-C3 alkoxy, (C1-C7 alkylamino)oxy, [di(C1-C7 alkyl)amino]oxy, (C3-C7 alkylideneamino)oxy, C1-C7 alkylamino, di(C1-C7 alkyl)amino, C3-C7 alkylylamino, C3-C7 alkynylamino, C5-C7 cycloalkylamino, [(C1-C7 alkoxy)carbonyl]C1-C3 alkylylamino, or (C1-C7 alkoxy)amino; R2 = H, or Me; R3 = H, halo, C1-C3 alkyly alkylamino, or (C1-C7 alkoxy)amino; R2 = H, or Me; R3 = H, branch carbon c

IT 380500-89-0P 477714-69-5P 477715-66-5

477715-68-7P

RL: AGR (Agricultural use); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation as plant growth regulator for cotton)

RN 380500-89-0 CAPLUS

Glycine, N-[[2-[2-chloro-5-[3,6-dihydro-3-methyl-2,6-dioxo-4-(trifluoromethyl)-1(2H)-pyrimidinyl]-4-fluorophenoxy]phenoxy]acetyl]-, methyl ester (9C1) (CA INDEX NAME)

RN 477714-69-5 CAPLUS

CN Alanine, N-[[2-[2-chloro-5-[3,6-dihydro-3-methyl-2,6-dioxo-4-(trifluoromethyl)-1(2H)-pyrimidinyl]-4-fluorophenoxy]phenoxy]acetyl]-2methyl-, methyl ester (9CI) (CA INDEX NAME)

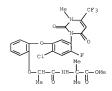
$$\begin{picture}(100,0) \put(0,0){\line(1,0){100}} \put(0,0){\line(1,0){10$$

RN 477715-66-5 CAPLUS

N Glycine, N-[2-[2-[2-chloro-5-[3,6-dihydro-3-methyl-2,6-dioxo-4-(trifluoromethyl)-1(2H)-pyrimidinyl]-4-fluorophenoxy]phenoxy]-1-oxopropyl]-, methyl ester (CA INDEX NAME)

RN 477715-68-7 CAPLUS

CN Alanine, N=[2-[2-[2-chloro-5-[3,6-dihydro-3-methyl-2,6-dioxo-4-(trifluoromethyl)-1(2H)-pyrimidinyl]-4-fluorophenoxy]phenoxy]-1-oxopropyl]-2-methyl-, methyl ester (CA INDEX NAME)



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 6 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2002:428894 CAPLUS Full-text

DOCUMENT NUMBER: 137:20303

TITLE: Preparation of substituted quinolines as antitumor

agents

INVENTOR(S): Boyle, Francis Thomas; Gibson, Keith Hopkinson; Foote, Kevin Michael

PATENT ASSIGNEE(S): Astrazeneca AB, Swed.; Astrazeneca UK Limited

SOURCE: PCT Int. Appl., 118 pp.

CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

						KIND DATE							DATE					
								2002	0000			2001				-	0011	026
	WO											2001-						
		W :										, BG,						
				), CR, CU, CZ, DE, DK, DM 1, HR, HU, ID, IL, IN, IS														
												, MW,						
	PL, PT, RO,								51,	SK,	, SL,	TJ,	TM,	TR,	TT,	TZ,	UA,	
	UG, US, UZ,																	
	RW: GH, GM, KE,																	
	DE, DK, ES,																	BF,
	BJ, CF, CG,																	
		U 2002010714															0011	
	EΡ	1337524				A1					EP 2	2001-	9786	16		2	0011	026
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	, IT,	LI,	LU,	NL,	SE,	MC,	PT,
			IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	, TR						
	JΡ	2004	5147	18		T		2004	0520		JP :	2002-		20011026				
	US	2004	0029	898		A1		2004	0212		US :	2003-		2	0030	502		
	US	7067	532			B2		2006	0627									
	US	2007	0021	407		A1		2007	0125	US 2006-374423						2	0060	314
	US 7402583							2008	0722									
PRIOR	RITY	APP	LN.	INFO	. :						GB 2	2000-	2674	4		A 2	0001	102
											GB 2	2000-	2674	6		A 2	0001	102
												2000-						
												2001-						
												2003-						
														~ ~				

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

- AB Title compds. I [n = 0 or 1; Y = NH, O, S, or alkylamine; R5 = CN, F, Cl, or Br; R6 = (un)substituted -cycloalkyl, -pyridinyl, -pyrimidinyl, -Ph, etc.; R1, R2 and R4 independently = H, OH, halo, CN, NO2, F3C, alkyl, amine, alkylamine, dialkylamine,  $R^{7}X1$  (CH2)x- wherein x = 0-3,  $R^{7}$  = H, (un)substituted hydrocarbyl or heterocyclyl and X1 = 0, CH2, OCO, CO, S, SO, SO2, NR8CO, NR8CO2, CONR9, CO2NR9, SO2NR10, NR11 or NR11NR11 wherein R8, R9, R10 and R11 independently = H, alkyl or alkoxyalkyl; R3 = group of formula X1R12(OH)p where p = 1-2 and R12 = alkylene, alkenylene or alkynylene chain, optionally interposed with a heteroatom or heterocyclic ring with the provision that when R12 = alkylene, R12 must be interposed with a heteroatom or heterocyclic ring and at least one (OH)p is on the alkylene chain between X1 and the interposed heteroatom or heterocyclic ring; group of formula R7(CH2) yX1(CH2)x where y = 0-5 and (CH2)y is optionally interposed by an X1 group; group of formula X1alkyl where alkyl is substituted by one or more Cl and/or CN; heterocyclic ring, etc.], or a pharmaceutically acceptable salt, pro-drug or solvate thereof are prepared and disclosed as antiproliferative agents. Thus, II was prepared in eight steps from benzylchloroformate and 2-methoxy-5-nitroaniline. I were evaluated as inhibitors of MAPK pathway and exhibited IC50 values typically lest than 0.5 μM, e.g., II possessed an IC50 = 0.0013μM. In cell proliferation assays, I had IC50 results typically less than  $30\mu M$  with II giving an IC50 of 1.3  $\mu M$  in HT29 human colon tumor cells. Methods for prevention of cancer comprising administering an effective amount of compound I are further claimed.
  - II 306999-95-1P 307309-82-6P
    RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
    (Reactant or reagent)

(intermediate; preparation, inhibition of MAP kinase, and cellular antiproliferation activity of substituted quinolines as antitumor agents)

- RN 306999-95-1 CAPLUS
- CN Glycine, N-[[2-(4-nitrophenoxy)phenoxy]acetyl]-, methyl ester (9CI) (CA INDEX NAME)

- RN 307309-82-6 CAPLUS
- CN Glycine, N-[[2-(4-aminophenoxy)phenoxy]acetyl]-, methyl ester (9CI) (CA INDEX NAME)

$$\text{MeO-} \overset{\circ}{\text{C-}} \text{CH}_2 - \text{NH} - \overset{\circ}{\text{C-}} \text{CH}_2 - \overset{\circ}{\text{O}}$$

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 7 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2002:353433 CAPLUS Full-text

DOCUMENT NUMBER: 136:369616

TITLE: Preparation of 3-cyano-4-arylaminoquinolines as inhibitors of MAP kinase for use as antitumor agents

INVENTOR(S): Boyle, Francis Thomas; Gibson, Keith Hopkinson

PATENT ASSIGNEE(S): Astrazeneca AB, Swed.; Astrazeneca UK Limited SOURCE: PCT Int. Appl., 149 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2 PATENT INFORMATION:

						KIND DATE													
		TENT :																DATE	
		2002																	
		W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA	, BI	В,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ	, E0	Ξ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JΡ	, KI	Ε,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK	, M	N,	ΜW,	MX,	MZ,	NO,	NZ,	PH,	PL,
	PT, RO, RU,						SE,	SG,	SI,	SK	, SI	L,	ΤJ,	TM,	TR,	TT,	TZ,	UA,	UG,
	US, UZ, VN,						ZA,	ZW											
		RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL	, S:	Z,	TZ,	UG,	ZW,	ΑT,	BE,	CH,	CY,
			DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE	, I:	Γ,	LU,	MC,	NL,	PT,	SE,	TR,	BF,
								GA,											
		U 2001095791																	
	ΕP	1337	513			A1 20030827					ΕP	20	01-	9765	23		2	20011	025
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB	, GI	R,	ΙT,	LI,	LU,	NL,	SE,	MC,	PT,
			ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY	, Al	L,	TR						
										JP 2002-539330						20011025			
		2005									US	20	03-	4158	13		- 2	20030	502
	US	7253	184			B2		2007	0807										
	US 20080027054										US	20	07-	8265	07		- 2	20070	716
	US 7504416							2009	0317										
PRIOR	IT:	Y APP	LN.	INFO	. :													20001	
											GB	20	00-	2674	7			20001	
															33			20011	
											US	20	03-	4158	13		A3 2	20030	502
OTHER	THER SOURCE(S):						PAT	136:369616											

<sup>\*</sup> STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Compds. I [R1, R2, R3, R4 independently H, HO, halogen, NC, O2N, F3C, (un) substituted C1-C3 alkyl, (un) substituted amino, saturated heterocyclyl containing two heteroatoms; R5 = NC, F, Cl, Br; R6 = divalent C1-C5 alkenyl, C3-C7 cycloalkyl, or heteroaryl moiety; R7 = AR8; A = bond, O, CO, S, SO, SO2, (un) substituted aminocarbonyl, (un) substituted carbonylamino, (un) substituted sulfonylamino, (un)substituted aminosulfonyl, (un)substituted amino; R8 = C1-C6 alky1, C2-C6 alkeny1, C2-C6 alkyny1; R9 = (un)substituted C3-C7 divalent cycloalkyl: R10 = (un)substituted arvlene, heteroarvlene, heteroarvlene Noxide, C3-C10 cycloalkylene; X = amino, (C1-C6)alkylamino, O, S, CH2; Y = amino, (C1-C6) alkylamino, O, S; Z = (un) substituted alkyl, alkylene, alkynylene, O, CO, COO, S, SO, SO2, (un)substituted aminocarbonyl, carbonylamino, sulfonylamino, aminosulfonyl, amino; n = 0,1; m and p independently 0-3; alternatively, R10Z(CH2)pR6R7 can be replaced with a heteroaryl or heterocyclyl-2,3-fused Ph ring] were prepared as inhibitors of MAP kinase for use as antitumor agents. E.g., 1-fluoro-4-nitrobenzene undergoes nucleophilic substitution with (2-hydroxyphenoxy)acetic acid followed by coupling of the acid with Me glycinate, reduction of the nitro group with Pd/C, and reaction of the ester with N-methylpiperazine to give the aminophenoxymethylcarbonylaminoacetyl piperazine II. E.g., coupling of II with 4-chloro-6,7-dimethoxy-3-quinolinenitrile gave the example compound III. Biol. data was obtained for selected compds. Selected compds. inhibited MAP kinase with IC50 < 0.5 μM; for example, III gave an IC50 of 3.8 nM. In addition, selected compds. inhibited the proliferation of human colon cancer cells with IC50 < 30 µM; for example, III gave an IC50 of 1 µM. 423179-57-1P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(example compds.; preparation of 4-arylamino-3-cyanoquinolines as inhibitors
of MAP kinase for potential use as antitumor agents)

RN 423179-57-1 CAPLUS

CN

Glycine, N-[[2-[4-[(3-cyano-6,7-dimethoxy-4-quinolinyl)amino]phenoxy]phenoxy]acetyl]-N-methyl-, methyl ester (9CI) (CA INDEX NAME)

IT 306999-95-1P 307309-82-6P 423180-30-7P

423180-31-6P 423180-57-8P 423180-59-0P

423180-89-6P 423180-90-9P 423180-96-5P 423180-97-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(intermediates; preparation of 4-arylamino-3-cyanoquinolines as inhibitors of MAP kinase for potential use as antitumor agents)

- RN 306999-95-1 CAPLUS
- CN Glycine, N-[[2-(4-nitrophenoxy)phenoxy]acetyl]-, methyl ester (9CI) (CA INDEX NAME)

- RN 307309-82-6 CAPLUS
- CN Glycine, N-[[2-(4-aminophenoxy)phenoxy]acetyl]-, methyl ester (9CI) (CA INDEX NAME)

- RN 423180-30-7 CAPLUS
- CN Glycine, N-[[2-(4-aminophenoxy)phenoxy]acetyl]-N-methyl-, ethyl ester (9CI) (CA INDEX NAME)

- RN 423180-31-8 CAPLUS
- CN Glycine, N-[[2-(4-aminophenoxy)phenoxy]acetyl]-N-methyl-, methyl ester (9CI) (CA INDEX NAME)

RN 423180-57-8 CAPLUS

CN Glycine, N-[2-[2-(4-aminophenoxy)phenoxy]-2-methyl-1-oxopropyl]-, ethyl ester (CA INDEX NAME)

RN 423180-59-0 CAPLUS

CN Glycine, N-[2-[2-[4-[(3-cyano-6,7-dimethoxy-4-quinoliny])amino]phenoxy]phenoxy]-2-methyl-1-oxopropyl]-, ethyl ester (CA INDEX NAME)

RN 423180-89-6 CAPLUS

CN Glycine, N-[difluoro[2-(4-nitrophenoxy)phenoxy]acetyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

RN 423180-90-9 CAPLUS

CN Glycine, N-[difluoro[2-(4-nitrophenoxy)phenoxy]acety1]- (9CI) (CA INDEX NAME)

RN 423180-96-5 CAPLUS

CN Glycine, N-[2-methyl-2-[2-(4-nitrophenoxy)phenoxy]-1-oxopropyl]-, ethyl ester (CA INDEX NAME)

RN 423180-97-6 CAPLUS

CN Glycine, N-[2-methyl-2-[2-(4-nitrophenoxy)phenoxy]-1-oxopropyl]- (CA INDEX NAME)

L3 ANSWER 8 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2002:171867 CAPLUS Full-text

DOCUMENT NUMBER: 136:232314

TITLE: Preparation of aminoquinazolines as epidermal growth factor receptor signal transduction inhibitors INVENTOR(S): Himmelsbach, Frank; Langkopf, Elke; Jung, Birgit;

Blech, Stefan; Solca, Flavio PATENT ASSIGNEE(S): Boehringer Ingelheim Pharma Kg, Germany

SOURCE: PCT Int. Appl., 103 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	TENT	NO.			KIND DATE					API	PLICA	DATE							
WO	2002		A1 20020307				WO	2001	20010818										
	W: AE, AG, AL		AL,	AM,	AT,	AU,	AZ,	BA,	BE	в, во	, BF	BY,	BZ,	CA,	CH,	CN,			
													, FI,						
													, KR,						
													, MZ,						
													1, TR,						
					YU,														
	RW:	GH,	GM,	KE,	LS,	MW.	MZ,	SD,	SL,	SZ	. T2	, UG	, ZW,	AT,	BE,	CH,	CY,		
													, NL,						
		BJ,	CF.	CG,	CI,	CM,	GA,	GN,	GO,	GV	, MI	, MF	, NE,	SN,	TD,	TG			
DE	1004	2058			A1		2002	0307		DE	2000	-100	42058		2	0000	826		
AU	2001	0876	94		A1 20020307 A 20020313					AU	2001	-876	94		20010818				
CA	2417	897			A1		2003	0130		CA	2001	-241	7897		20010818				
EP	1315	705			A1	2003	0604		EP	2001	-967	285		20010818					
	R:												, LU,						
		IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AI	, TE								
BR	2001	0135	19		A		2003	0701		BR	2001	-135	19		2	0010	818		
HU	2003	19		A2		BR 2001-13519 HU 2003-819 JP 2002-523469 EE 2003-77 NZ 2001-524668 AU 2001-287694 IL 2001-154602 CN 2001-814635						0010	818						
HU	2003	0008	19		A3		2008	0328											
JP	2004	5075	29		T		JP	2002	-523	469		2	0010	818					
EE	2003	0007	7		A		EE	2003	-77			20010818							
NZ	5246	68			Α		NZ	2001		20010818									
AU	2001	2876	94		B2		AU	2001		20010818									
IL	1546	02			Α		IL	2001		20010818									
CN	1004	0451	7		C		CN	2001	-814		20010818								
US	2002	0082	271		A1		US	2001	-934		20010822								
US	6656	946			B2		2003	1202											
ZA	2003	0009	91		A		2004	0416		$z_{A}$	2003	-991			2	0030	205		
BG	1075	59			A		2003		ZA 2003-991 BG 2003-107559 IN 2003-MN222 MY 2003-1483					2	0030	214			
IN	2003	MNOO	222		A		2005	0211		IN	N 2003-MN222				2	0030	214		
NO	2003	0008	70		A		2003	0225		NO	2003	-870	)		2	0030	225		
NO	3248	66			B1		2007	1217											
KR	NO 2003000870 NO 324866 KR 862873						2008	1015		KR	2003	-702	744		2	0030	225		
HK	1057	557			A1		2008	1031		HK	2004	-100		20040121					
PRIORIT																A 20000826			
										US	2000	-230	035P		P 2	0000	905		
										WO	2001	-EP9	532		W 2	0010	818		
OTHER C	OUDOR	M2 D1	D 7 T	126.	2222	1.4													

- Title compds. [I; R1 = PhCH2, 1-phenylethyl, (substituted) Ph; R2, R3 = AB O(CH2) mR4, methoxy, cyclobutyloxy, cyclopentyloxy, cyclopropylmethoxy, cyclobutylmethoxy, cyclopentylmethoxy, tetrahydrofuran-3-yloxy, tetrahydropyran-3-vloxy, tetrahydropyran-4-vloxy, tetrahydrofuranylmethoxy, tetrahydropyranylmethoxy; R4 = N-(2-oxotetrahydrofuran-4-yl)methylamino, N-(2oxotetrahydrofuran-4-vl)ethylamino, (substituted) 2-oxo-morpholin-4-vl, R50C0CH2NCH2CH2OH; R5 = H, alkyl; m = 2-4], were prepared Thus, 4-[(3-4)]bromophenyl)aminol-6-[2-(N-[(tert-butyloxycarbonyl)methyl]-N-((S)-2hydroxypropyl)amino)ethoxy]-7- methoxyquinazoline (preparation given) in MeCN was stirred under reflux with MeSO2OH for 3 h followed by addition of MeSO2OH up to completely conversion to give 85% 4-[(3-bromophenyl)amino]-6-[2-((S)-6methyl-2-oxomorpholin-4- yl)ethoxy]-7-methoxyquinoline. Tested I inhibited epidermal growth factor (EGF)-dependent proliferation of F/L-HERc cells with IC50 = 29-59 nM. The invention relates to the use of the title compds. for treating tumor diseases, and lung and respiratory tract disorders.
  - 402735-26-6P 402735-27-7P 402735-34-6P

402735-35-7P 402735-36-8P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of aminoquinazolines as epidermal growth factor receptor signal transduction inhibitors)  $\begin{tabular}{ll} \hline \end{tabular}$ 

RN 402735-26-6 CAPLUS

CN Glycine, N-[2-[[4-[(3-chloro-4-fluorophenyl)amino]-7-(cyclopentyloxy)-6-quinazolinyl)axylethyl]-N-[(25)-2-hydroxypropyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

Absolute stereochemistry.

- RN 402735-27-7 CAPLUS
- CN Glycine, N-[2-[[4-[(3-chloro-4-fluoropheny1)amino]-7-(cyclopentyloxy)-6quinazolinyl]oxy]ethyl]-N-[(2R)-2-hydroxypropyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

Absolute stereochemistry.

RN 402735-34-6 CAPLUS

CN Glycine, N-[2-[[4-[(3-chloro-4-fluorophenyl)amino]-7-(cyclopentyloxy)-6quinazolinyl]oxy]ethyl]-N-(2-hydroxy-2-methylpropyl)-, ethyl ester (CA INDEX NAME)

RN 402735-35-7 CAPLUS

CN Glycine, N-[2-[[4-[(3-chloro-4-fluorophenyl)amino]-6-(cyclopentyloxy)-7quinazolinyl]oxy|sthyl]-N-[(28)-2-hydroxypropyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

Absolute stereochemistry.

RN 402735-36-8 CAPLUS

CN Glycine, N-[2-[[4-[(3-chloro-4-fluorophenyl)amino]-6-(cyclopentyloxy)-7quinazolinyl]oxy]ethyl]-N-[(2R)-2-hydroxypropyl]-, 1,1-dimethylethyl ester (CA INDEX INDEX INME)

Absolute stereochemistry.

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 9 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2001:910259 CAPLUS Full-text

DOCUMENT NUMBER: 136:53754

TITLE: Preparation and application of uracils as herbicides

INVENTOR(S): Goto, Tomohiko; Sanemitsu, Minoru PATENT ASSIGNEE(S): Sumitomo Chemical Co., Ltd., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 91 pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent

LANGUAGE: Japanese FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DDT	JP 2001348376 DRITY APPLN. INFO.:	A	20011218	JP 2000-170234 JP 2000-170234	20000607
	ER SOURCE(S):	MARPAT	136:53754	JP 2000-170234	20000607
GI					

- AB Title compds. [I; R = OCH(CH3)COOCH2COOCH3, (S)-OCH2CONHCH(CH2CH(CH3)2)CO2CH3, OCH2CONHCH2CO2CH3, OCH2CO2CH3, OCH2CO2CH3, OCH2CO2CH3, OCH2CO2CH3, OCH2CO2CH2CH2CH2CH2, H, CF3, CH3; R1 = H, OCH(CH3)CO2CH2COOCH, OCH2COOCH2COOCH2CH:CH2, H; R2 = H, OCH(CH3)CO2CH2COOCH, OCH2COOCH2COSCH2CH3; X = F, H; Y = Cl, NO2; X1 = O, S, NH] are prepared as herbicides. Thus, the title compound I (R = OCH2COOC (CH3)2COOCH2CH2CH2CH2; R1 = H; R2 = H; X = F; X1 = O; Y = Cl) was prepared and field tested as effective herbicide in forage and soil treatment. IT 38050-89-07 28050-90-39
  - RL: AGR (Agricultural use); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation and application of uracils as herbicides)

RN 380500-89-0 CAPLUS

CN Glycine, N-[[2-[2-chloro-5-[3,6-dihydro-3-methyl-2,6-dioxo-4-(trifluoromethyl)-1(2H)-pyrimidinyl]-4-fluorophenoxy]phenoxy]acetyl]-, methyl ester (9C1) (CA INDEX NAME)

RN 380500-90-3 CAPLUS

IN L-Leucine, N-[12-[2-chloro-5-[3,6-dihydro-3-methyl-2,6-dioxo-4-(trifluoromethyl)-1(2H)-pyrimidinyl]-4-fluorophenoxy]phenoxy]acetyl]-, methyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L3 ANSWER 10 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2000:814464 CAPLUS Full-text

DOCUMENT NUMBER: 133:362712

TITLE: Preparation of quinoline derivatives as inhibitors of

MEK enzymes

INVENTOR(S): Boyle, Francis Thomas; Gibson, Keith Hopkinson;

Poyser, Jeffrey Philip; Turner, Paul

PATENT ASSIGNEE(S): Astrazeneca AB, Swed.

SOURCE: PCT Int. Appl., 187 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.						KIND DATE											
	2000 W:	AE, CU, ID, LV, SG, GH,	O1 AG, CZ, IL, MA, SI, GM,	AL, DE, IN, MD, SK, KE,	A1 AM, DK, IS, MG, SL, LS,	AT DM JP MK TJ MW		AZ, EE, KG, MW, TR, SL,	BA, ES, KP, MX, TT,	WO 2 BB, FI, KR, NO, TZ,	2000- BG, BB, KZ, NZ, UA,	GB16 BR, GD, LC, PL, UG, ZW,	97 BY, GE, LK, PT, US, AT,	CA, GH, LR, RO, UZ, BE,	CH, GM, LS, RU, VN, CH,	CN, HR, LT, SD, YU, CY,	CR, HU, LU, SE, ZA, DE,	ZW
EP	2372 1178 1178	CG, 663 9 <b>6</b> 7	CI,	CM,	GA, A1 A1	GN	GW, 2000	ML, 1116 0213	MR,	NE,	, SN, 2000-	TD, 2372	TG 663		2	0000	503	
	R:	AT, IE,	BE, SI,	LT,	DE, LV,	DK FI	ES,	FR, CY	GB,									
BR HU	2000010391				A A2	2002	0702 0928		TR 2001-3186 BR 2000-10391 HU 2002-1219						20000503			
EE NZ	2002001219 200100589 514980				A A		2003	0217		EE 2 NZ 2	2001- 2000-	589 5149		20000503 20000503 20000503				
CN	1219 1584	768 619			C A1		2005 2005	0921 1012		CN 2 EP 2	2000- 2005-	8099 1358	59 7		2	0000 0000	503 503	
AT	3196	IE,	LV,	FI,	MK,	CY	2006	0315	·	АТ 2	2000-	9274	91	·	2	0000	503	
PT ES ZA	1178 2258 2001	967 455 0089	71		T T3 A		2006 2006 2003	0630 0901 0130		PT 2 ES 2 ZA 2	2000- 2000- 2001-	9274 9274 8971	91 91		2 2 2	0000 0011	503 030	
BG NO	NO 2001005448					A 20020531 A 20020107					BG 2001-106073 NO 2001-5448					20011101		
NO MX	NO 321696				B1		2006 2002	0626 0311		MX 2001-11360 GB 1999-10577 EP 2000-927491					A 19990508			
OTHER SO	OURCE	(S):			MARI	PAT	133:	3627			2000-							

<sup>\*</sup> STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Title compds. [I; or a pharmaceutically acceptable salt thereof wherein: n is 0-1; X and Y are independently selected from NH, O, S, or NR8 where R8 is alkyl of 1-6 carbon atoms and X may addnl. comprise a CH2 group; R7 is a group (CH2)mR9 where m is 0, or an integer of from 1-3 and R9 is a substituted aryl group, an optionally substituted cycloalkyl ring of up to 10 carbon atoms, or an optionally substituted heterocyclic ring or an N-oxide of any nitrogen containing ring; R6 is a divalent cycloalkyl of 3 to 7 carbon atoms, which may be optionally further substituted with one or more alkyl of 1 to 6 carbon atom group; or is a divalent cycloalkyl pyrimdidnyl, or Ph ring; wherein the

pyridinyl, pyrimidinyl, or Ph ring may be optionally further substituted with one or more specified groups; Rl, R2, R3 and R4 are each independently selected from hydrogen or various specified organic groups]. Title compous are useful as pharmaceuticals for the inhibition of MEK activity. Thus, the title compound II was prepared and tested in HT29 human colon tumor cell proliferation assay.

IT 306999-63-3P 306999-65-5P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation of quinoline derivs. as inhibitors of MEK enzymes)

RN 306999-63-3 CAPLUS

CN L-Glutamic acid, N-[[2-[4-[(3-cyano-6,7-dimethoxy-4-quinoliny1)amino]phenoxy]phenoxy]acety1]-, dimethyl ester (9CI) (CA INDEX NAME)

## Absolute stereochemistry.

RN 306999-65-5 CAPLUS

CN Glycine, N-[[2-[4-[(3-cyano-6,7-dimethoxy-4-quinoliny1)amino]phenoxy]phenoxy]acetyl]-, methyl ester (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{MeO} \\ \text{MeO} \\ \end{array}$$

IT 306999-81-5 306999-85-9

RL: RCT (Reactant); RACT (Reactant or reagent) (preparation of quinoline derivs. as inhibitors of MEK enzymes)

RN 306999-81-5 CAPLUS

CN L-Glutamic acid, N-[[2-(4-aminophenoxy)phenoxy]acetyl]-, dimethyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 306999-85-9 CAPLUS

CN L-Alanine, N-[[2-(4-aminophenoxy)phenoxy]acetyl]-, ethyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

- IT 306999-93-9P 306999-95-1P 306999-96-2P 307309-82-6P
  - RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
  - (preparation of quinoline derivs. as inhibitors of MEK enzymes)
- RN 306999-93-9 CAPLUS
- CN L-Glutamic acid, N-[[2-(4-nitrophenoxy)phenoxy]acetyl]-, dimethyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 306999-95-1 CAPLUS

CN Glycine, N-[[2-(4-nitrophenoxy)phenoxy]acetyl]-, methyl ester (9CI) (CA

INDEX NAME)

RN 306999-96-2 CAPLUS

CN L-Alanine, N-[[2-(4-nitrophenoxy)phenoxy]acetyl]-, ethyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 307309-82-6 CAPLUS

CN Glycine, N-[[2-(4-aminophenoxy)phenoxy]acetyl]-, methyl ester (9CI) (CA INDEX NAME)

REFERENCE COUNT:

3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 11 OF 11 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2000:666715 CAPLUS Full-text DOCUMENT NUMBER: 133:252449

TITLE: Quinazolines and other bicyclic heterocycles,

pharmaceutical compositions containing these compounds as tyrosine kinase inhibitors, and processes for preparing them

INVENTOR(S): Himmelsbach, Frank; Langkopf, Elke; Blech, Stefan;

Jung, Birgit; Metz, Thomas; Solca, Flavio
PATENT ASSIGNEE(S): Boehringer Ingelheim Pharma K.-G., Germany

SOURCE: PCT Int. Appl., 153 pp.

## CODEN: PIXXD2

Patent English

DOCUMENT TYPE: FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

LANGUAGE:

PA'	TENT	NO.			KIND DATE					API	PL:	ICAT:	DATE								
	2000						2000						20000314								
	W:	ΑE,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BO	3,	BR,	BY,	CA,	CH,	CN,	CR,	CU,			
		CZ,	DE,	DK,	DM,	EE,	ES,	FI,	GB,	GI	ο,	GE,	GH,	GM,	HR,	HU,	ID,	IL,			
		IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LO	Ξ,	LK,	LR,	LS,	LT,	LU,	LV,	MA,			
		MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PI	Ŀ,	PT,	RO,	RU,	SD,	SE,	SG,	SI,			
																	, ZW				
	RW:																				
															SE,	BF,	BJ,	CF,			
				CM,	GA,	GN,	GW,	ML,	MR,	NE	Ξ,	SN,	TD,	TG							
	1991				A1		2000	0921		DE	19	999-:	1991	1509		19990315 20000314					
	2368		A1		2000	0921		CA	20	000-2	2368	059		20000314							
	1163				A1		2001	1219		EP	20	000-9	9093	60		20000314					
EP	1163									R, IT, LI, LU,											
	R:							FR,	GB,	GE	٦,	IT,	LI,	LU,	NL,	SE,	MC,	PT,			
					LV,	FI,	RO														
BR	2000	0090	/6		A		2001	1226		BR	20	000-9	9076			- 1	20000	314			
TR	2001	0278	2		A 20011226 T2 20020422					TR	20	001-2	2/82			20000314 20000314					
JP	2002	2391	99		T		2002			JP 2000-605571						20000314					
JP	3/54	0010	4		B2 20060315 A 20021216					EE 2001-484						20000314					
22	E034	0040	*		D1	22 2001-404							20000314								
DIT DE	200102782 2002539199 3754617 200100484 5034 2002001832 2002001832				V 3		2008			HU 2002-1832							20000	91/			
HII	2002	32		7.3	A2 20021228 HU 2002-1832 A3 20030228								20000314								
NZ.	5147	06 TO.	22		A		2003			NZ 2000-514706							20000	314			
AII	7725	20			B2		2004			AU 2000-31667						- 3	20000	314			
CN	1150	171			C		2004	0519		AU 2000-31667 CN 2000-805005						20000314					
AT	3054	56			A2 20021228 A3 20030228 A 20031128 B2 20040429 C 20040519 T 20051015 T3 20060416 A 20070211 B 20061221 A 20050304 A 20020314					AT	20	000-9	9093		20000314						
ES	2250	111			Т3		2006	0416		AT 2000-909360 ES 2000-909360 IL 2000-144626							20000314				
IL	1446	26			A	0211		IL	20	000-	1446	20000314									
TW	2689	24			В		TW	20	000-8	8910	20000426 20010809										
IN	2001	MNOO	956		A		2005	0304		IN	20	001-1	MN95	20010809							
MX	2001	0083	24		A		2002	0311		MΧ	20	2001-MN956 2001-8324 2001-938235 2001-7185 2001-105893					20010816				
US	2002	0177	601		A1		2002	1128	US 2001-938235						20010823						
ZA	2001	0071	85		A		2002	20020621			ZA 2001-7185				20010830						
BG	1058	93			A		2002	0531		BG	20	001-	1058	93		- 2	20010	912			
BG	6513	0			B1		2007	0330													
KR	7492			B1		2007	20020311 20021128 20020621 20020531 20070330 20070814 20010914 200409123			20	001-	1-711645 1-4487 2-104697			- 2	20010	913				
ИО	2001	В7		A		2001	0914		ИО	20	001-	4487			- 2	20010	914				
HK	1043	124			A1		2004	1203		HK	20	002-:	1046	97		20020625					
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The invention relates to bicyclic heterocyclic compds. I [R1 = H, alkyl; R2 = AB (un) substituted Ph, CH2Ph, or CH(Me)Ph; R3, R4 = H, F, C1, OMe, or Me optionally substituted by OMe, NMe2, NEt2, pyrrolidino, piperidino, or morpholino; X = N or C(CN); A = O, NH, (un)substituted alkylene, O-alkylene, NH-alkylene, O-cycloalkylene, etc.; B = (un) substituted amine-containing sidechain, piperazino, alkyleneimino, morpholino, etc.; or AB = H, F, Cl, alkoxy, amino, etc.; C = groups similar to A; D = groups similar to B; with a variety of provisos | and their tautomers, stereoisomers, and salts, and particularly their physiol, acceptable salts with inorg, or organic acids or bases. The compds, have valuable pharmacol, properties, particularly an inhibitory effect on signal transduction mediated by tyrosine kinases, and are useful in treating diseases, particularly tumor diseases, and diseases of the lung and airways. Over 20 compds. were prepared, and over 200 are listed. For instance, alkylation of 4-(3-chloro-4-fluorophenylamino)-6-[3-(1piperazinyl)propyloxyl-7- methoxyquinazoline (preparation given) by Me bromoacetate gave 51% title compound II. The latter compound inhibited EGFdependent proliferation of F/L-HERC cells in vitro, with an IC50 of 46 nM. ΙT 295330-29-9P, 4-[(3-Chloro-4-fluorophenyl)amino]-6-cyclopentyloxy-

7-[2-[N-(2-hydroxy-2-methylprop-1-yl)-N-

[(ethoxycarbonyl)methyl]amino]ethoxy]quinazoline
RL: BAC (Biological activity or effector, except adverse); BSU (Biological
study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of quinazoline derivs. and other bicyclic heterocycles as tyrosine kinase inhibitors)

RN 295330-29-9 CAPLUS

CN Glycine, N-[2-[[4-[(3-chloro-4-fluoropheny])amino]-6-(cyclopentyloxy)-7-quinazolinyl]oxy]ethyl]-N-(2-hydroxy-2-methylpropyl)-, ethyl ester (CA INDEX NAME)

REFERENCE COUNT: 5

THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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